

WHAT IS CLAIMED IS:

1. A substantially pure τ -conotoxin peptide selected from the group consisting of:

Phe-Cys-Cys-Xaa₁-Val-Ile-Arg-Xaa₂-Cys-Cys-Xaa₃ (SEQ ID NO:2);

Cys-Cys-Gln-Thr-Phe-Xaa₂-Xaa₃-Cys-Cys-Gln (SEQ ID NO:4);

Xaa₄-Gly-Xaa₃-Cys-Cys-Xaa₅-Xaa₆-Asn-Ile-Ala-Cys-Cys-Ile (SEQ ID NO:5);

Gly-Cys-Cys-Ala-Arg-Leu-Thr-Cys-Cys-Val (SEQ ID NO:6);

Asn-Gly-Cys-Cys-Xaa₁-Xaa₅-Gln-Met-Arg-Cys-Cys-Thr (SEQ ID NO:7);

Asp-Xaa₃-Asn-Ser-Cys-Cys-Gly-Xaa₅-Asn-Xaa₁-Gly-Cys-Cys-Xaa₁-Xaa₃ (SEQ ID NO:8);

Xaa₄-Gly-Xaa₃-Cys-Cys-Xaa₅-Xaa₆-Asn-Ile-Arg-Cys-Cys-Val (SEQ ID NO:9);

Xaa₆-Cys-Cys-Xaa₆-Asp-Gly-Xaa₃-Cys-Cys-Thr-Ala-Ala-Xaa₁-Leu-Thr (SEQ ID NO:10);

Gly-Cys-Cys-Xaa₆-Asp-Gly-Xaa₃-Cys-Cys-Thr-Ala-Ala-Xaa₁-Leu-Thr (SEQ ID NO:11);

Asn-Gly-Cys-Cys-Arg-Ala-Gly-Asp-Cys-Cys-Ser-Arg-Phe-Xaa₆-Ile-Xaa₅-Xaa₆-Asn-Asp-Phe (SEQ ID NO:12);

Asn-Ala-Cys-Cys-Ile-Val-Arg-Gln-Cys-Cys (SEQ ID NO:13);

Asn-Gly-Cys-Cys-Arg-Ala-Gly-Asp-Cys-Cys-Ser (SEQ ID NO:14);

Cys-Cys-Xaa₁-Arg-Arg-Leu-Ala-Cys-Cys-Ile-Ile (SEQ ID NO:15);

Cys-Cys-Xaa₁-Asn-Xaa₅-Xaa₁-Cys-Cys-Phe-Ile (SEQ ID NO:16);

Gly-Cys-Cys-Ala-Met-Leu-Thr-Cys-Cys-Val (SEQ ID NO:17);

Leu-Cys-Cys-Val-Thr-Xaa₆-Asp-Xaa₃-Cys-Cys-Xaa₆-Xaa₃-Xaa₃ (SEQ ID NO:18); and

Val-Cys-Cys-Arg-Xaa₁-Val-Gln-Asp-Cys-Cys-Ser (SEQ ID NO:19);

wherein Xaa₁ is Pro or hydroxy-Pro; Xaa₂ is Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr or nitro-Tyr; Xaa₃ is Trp or halo-Trp; Xaa₄ is Gln or pyro-Glu; Xaa₅ is Lys, N-methyl-Lys, N,N-dimethyl-Lys or N,n,N-trimethyl-Lys, Xaa₆ is Glu or gamma-carboxy-Glu (Gla); and the C-terminus contains a carboxyl or amide group.

2. The substantially pure τ -conotoxin peptide of claim 1, wherein Xaa₆ is Glu.
3. The substantially pure τ -conotoxin peptide of claim 1, wherein Xaa₅ is Lys.
4. The substantially pure τ -conotoxin peptide of claim 1, wherein Xaa₄ is Gln.
5. The substantially pure τ -conotoxin peptide of claim 1, wherein Xaa₂ is mono-iodo-Tyr.
6. The substantially pure τ -conotoxin peptide of claim 1, wherein Xaa₂ is di-iodo-Tyr.
7. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:2, wherein Xaa₁ is Pro, Xaa₂ is Tyr and Xaa₃ is Trp.

8. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:4, wherein Xaa₂ is Tyr and Xaa₃ is Trp.
9. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:5, wherein wherein Xaa₃ is Trp, Xaa₄ is Gln, Xaa₅ is Lys and Xaa₆ is Glu.
10. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:6.
11. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:7, wherein Xaa₁ is Pro and Xaa₅ is Lys.
12. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:8, wherein Xaa₁ is Pro, Xaa₃ is Trp and Xaa₅ is Lys.
13. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:9, wherein Xaa₃ is Trp, Xaa₄ is Gln, Xaa₅ is Lys and Xaa₆ is Glu.
14. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:10, wherein Xaa₁ is Pro, Xaa₃ is Trp and Xaa₆ is Glu.

15. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:11, wherein Xaa₁ is Pro, Xaa₃ is Trp and Xaa₆ is Glu.
16. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:12, wherein Xaa₅ is Lys and Xaa₆ is Glu.
17. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:13.
18. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:14.
19. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:15, wherein Xaa₁ is Pro.
20. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:16, wherein Xaa₁ is Pro and Xaa₅ is Lys.
21. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:17.
22. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:18, wherein Xaa₃ is Trp and Xaa₆ is Glu.

23. The substantially pure τ -conotoxin peptide of claim 1, wherein said peptide has the sequence set forth in SEQ ID NO:19, wherein Xaa₁ is Pro.
24. A method for inducing analgesia in an individual which comprises administering an effective amount of a substantially pure τ -conotoxin peptide having the generic formula I: Xaa₁-Xaa₂-Xaa₃-Xaa₄-Cys-Cys-Xaa₅-Xaa₆-Xaa₇-Xaa₈-Xaa₉-Cys-Cys-Xaa₁₀-Xaa₁₁-Xaa₁₂-Xaa₁₃-Xaa₁₄-Xaa₁₅-Xaa₁₆-Xaa₁₇-Xaa₁₈-Xaa₁₉ (SEQ ID NO:1), wherein Xaa₁ is des-Xaa₁, Asp, Glu or γ -carboxy-Glu (Gla); Xaa₂ is des-Xaa₂, Gln, Asn, Glu, Trp (D or L), neo-Trp, halo-Trp or any unnatural aromatic amino acid; Xaa₃ is des-Xaa₃, Gly, Ala, Asn or Gln; Xaa₄ is des-Xaa₄, Val, Leu (D or L), Ile, Ala, Gly, Glu, Gla, Asp, Ser, Thr, Phe, Trp (D or L), neo-Trp, halo-Trp (D or L) or any unnatural aromatic amino acid; Xaa₅ is Pro, hydroxy-Pro, Gln, Asn, Glu, Gla, Ala, Gly, Lys, Arg, Ile, Val, homoarginine, ornithine, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any unnatural basic amino acid; Xaa₆ is Val, Phe, Thr, Ser, Glu, Gla, Asp, Asn, Gln, Ala, Gly, Ile, Leu (D or L), Met, Pro, hydroxy-Pro, Arg, homoarginine, ornithine, Lys, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, any unnatural basic amino acid or any unnatural aromatic amino acid; Xaa₇ is any Val, Ile, Asn, Leu (D or L), Gln, Gly, Ala, Phe, Glu, Gla, Arg, ornithine, homoarginine, Lys, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, any unnatural basic amino acid or any unnatural aromatic amino acid; Xaa₈ is Ile, Leu (D or L), Met, Thr, Ser, Pro, hydroxy-Pro, Gln, Asp, Glu, Gla, Asn, Arg, homoarginine, ornithine, Lys, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr,

O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr, any unnatural basic amino acid, any unnatural aromatic amino acid or any unnatural hydroxy containing amino acid; Xaa₉ is des-Xaa₉, Ala, Gly, Asp, Glu, Glu, Trp (D or L) neo-Trp, halo-Trp (D or L), Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, Arg, homoarginine, ornithine, Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr or any unnatural basic amino acid; Xaa₁₀ is des-Xaa₁₀, Ile, Leu (D or L), Val, Glu, Glu, Asp, Thr, Ser, Pro, hydroxy-Pro, Trp (D or L), neo-Trp, halo-Trp (D or L), Phe, any unnatural aromatic amino acid or any unnatural hydroxy containing amino acid; Xaa₁₁ is des-Xaa₁₁, Gln, Asn, Leu (D or L), Ile, Val, Ala, Gly, Trp (D or L), neo-Trp, halo-Trp (D or L), Arg, homoarginine, ornithine, Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, any unnatural basic amino acid or any unnatural aromatic amino acid; Xaa₁₂ is des-Xaa₁₂, Ala, Gly, Phe, Trp (D or L), neo-Trp, halo-Trp (D or L) or any unnatural aromatic amino acid; Xaa₁₃ is des-Xaa₁₃, Glu, Glu, Asp, Phe or any unnatural aromatic amino acid; Xaa₁₄ is des-Xaa₁₄, Ile, Val or Leu (D or L); Xaa₁₅ is des-Xaa₁₅, Thr, Ser, Arg, homoarginine, ornithine, Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any unnatural basic amino acid; Xaa₁₆ is des-Xaa₁₆, Glu, Glu or Asp; Xaa₁₇ is des-Xaa₁₇, Asn or Gln; Xaa₁₈ is des-Xaa₁₈, Asp, Glu or Glu; Xaa₁₉ is des-Xaa₁₉, Phe or any unnatural aromatic amino acid; and the C-terminus contains a free carboxyl group or an amide group.

25. A method for inducing analgesia in an individual which comprises administering an effective amount of the τ -conotoxin peptide of claim 1.

26. The method of claim 24, wherein the substantially pure τ -conotoxin peptide is modified to contain an O-glycan, an S-glycan or an N-glycan.
27. The method of claim 25, wherein the substantially pure τ -conotoxin peptide is modified to contain an O-glycan, an S-glycan or an N-glycan.
28. A method for inducing analgesia in an individual which comprises administering an effective amount of a pharmaceutical composition comprising a τ -conotoxin peptide or a pharmaceutically acceptable salt thereof and a pharmaceutically acceptable carrier, said τ -conotoxin peptide having the generic formula I: Xaa₁-Xaa₂-Xaa₃-Xaa₄-Cys-Cys-Xaa₅-Xaa₆-Xaa₇-Xaa₈-Xaa₉-Cys-Cys-Xaa₁₀-Xaa₁₁-Xaa₁₂-Xaa₁₃-Xaa₁₄-Xaa₁₅-Xaa₁₆-Xaa₁₇-Xaa₁₈-Xaa₁₉ (SEQ ID NO:1), wherein Xaa₁ is des-Xaa₁, Asp, Glu or γ -carboxy-Glu (Gla); Xaa₂ is des-Xaa₂, Gln, Asn, Glu, Trp (D or L), neo-Trp, halo-Trp or any unnatural aromatic amino acid; Xaa₃ is des-Xaa₃, Gly, Ala, Asn or Gln; Xaa₄ is des-Xaa₄, Val, Leu (D or L), Ile, Ala, Gly, Glu, Gla, Asp, Ser, Thr, Phe, Trp (D or L), neo-Trp, halo-Trp (D or L) or any unnatural aromatic amino acid; Xaa₅ is Pro, hydroxy-Pro, Gln, Asn, Glu, Gla, Ala, Gly, Lys, Arg, Ile, Val, homoarginine, ornithine, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any unnatural basic amino acid; Xaa₆ is Val, Phe, Thr, Ser, Glu, Gla, Asp, Asn, Gln, Ala, Gly, Ile, Leu (D or L), Met, Pro, hydroxy-Pro, Arg, homoarginine, ornithine, Lys, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, any unnatural basic amino acid or any unnatural aromatic amino acid; Xaa₇ is any Val,

Ile, Asn, Leu (D or L), Gln, Gly, Ala, Phe, Glu, Gla, Arg, ornithine, homoarginine, Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, any unnatural basic amino acid or any unnatural aromatic amino acid; Xaa₈ is Ile, Leu (D or L), Met, Thr, Ser, Pro, hydroxy-Pro, Gln, Asp, Glu, Gla, Asn, Arg, homoarginine, ornithine, Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr, any unnatural basic amino acid, any unnatural aromatic amino acid or any unnatural hydroxy containing amino acid; Xaa₉ is des-Xaa₉, Ala, Gly, Asp, Glu, Gla, Trp (D or L) neo-Trp, halo-Trp (D or L), Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, Arg, homoarginine, ornithine, Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr or any unnatural basic amino acid; Xaa₁₀ is des-Xaa₁₀, Ile, Leu (D or L), Val, Glu, Gla, Asp, Thr, Ser, Pro, hydroxy-Pro, Trp (D or L), neo-Trp, halo-Trp (D or L), Phe, any unnatural aromatic amino acid or any unnatural hydroxy containing amino acid; Xaa₁₁ is des-Xaa₁₁, Gln, Asn, Leu (D or L), Ile, Val, Ala, Gly, Trp (D or L), neo-Trp, halo-Trp (D or L), Arg, homoarginine, ornithine, Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys, any unnatural basic amino acid or any unnatural aromatic amino acid; Xaa₁₂ is des-Xaa₁₂, Ala, Gly, Phe, Trp (D or L), neo-Trp, halo-Trp (D or L) or any unnatural aromatic amino acid; Xaa₁₃ is des-Xaa₁₃, Glu, Gla, Asp, Phe or any unnatural aromatic amino acid; Xaa₁₄ is des-Xaa₁₄, Ile, Val or Leu (D or L); Xaa₁₅ is des-Xaa₁₅, Thr, Ser, Arg, homoarginine, ornithine, Lys, N-methy-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any unnatural basic amino acid; Xaa₁₆ is des-Xaa₁₆, Glu, Gla or Asp; Xaa₁₇ is des-Xaa₁₇, Asn or Gln;

Xaa₁₈ is des-Xaa₁₈, Asp, Glu or Gla; Xaa₁₉ is des-Xaa₁₉, Phe or any unnatural aromatic amino acid; and the C-terminus contains a free carboxyl group or an amide group.

29. The method of claim 28, wherein the pharmaceutical composition is modified to contain an O-glycan, an S-glycan or an N-glycan.

30. A method for inducing analgesia in an individual which comprises administering an effective amount of a pharmaceutical composition comprising a τ -conotoxin peptide or a pharmaceutically acceptable salt thereof and a pharmaceutically acceptable carrier, said τ -conotoxin peptide selected from the group consisting of:

Phe-Cys-Cys-Xaa₁-Val-Ile-Arg-Xaa₂-Cys-Cys-Xaa₃ (SEQ ID NO:2);

Phe-Cys-Cys-Xaa₁-Phe-Ile-Arg-Xaa₂-Cys-Cys-Xaa₃ (SEQ ID NO:3);

Cys-Cys-Gln-Thr-Phe-Xaa₂-Xaa₃-Cys-Cys-Gln (SEQ ID NO:4);

Xaa₄-Gly-Xaa₃-Cys-Cys-Xaa₅-Xaa₆-Asn-Ile-Ala-Cys-Cys-Ile (SEQ ID NO:5);

Gly-Cys-Cys-Ala-Arg-Leu-Thr-Cys-Cys-Val (SEQ ID NO:6);

Asn-Gly-Cys-Cys-Xaa₁-Xaa₅-Gln-Met-Arg-Cys-Cys-Thr (SEQ ID NO:7);

Asp-Xaa₃-Asn-Ser-Cys-Cys-Gly-Xaa₅-Asn-Xaa₁-Gly-Cys-Cys-Xaa₁-Xaa₃ (SEQ ID NO:8);

Xaa₄-Gly-Xaa₃-Cys-Cys-Xaa₅-Xaa₆-Asn-Ile-Arg-Cys-Cys-Val (SEQ ID NO:9);

Xaa₆-Cys-Cys-Xaa₆-Asp-Gly-Xaa₃-Cys-Cys-Thr-Ala-Ala-Xaa₁-Leu-Thr (SEQ ID NO:10);

Gly-Cys-Cys-Xaa₆-Asp-Gly-Xaa₃-Cys-Cys-Thr-Ala-Ala-Xaa₁-Leu-Thr (SEQ ID NO:11);

Asn-Gly-Cys-Cys-Arg-Ala-Gly-Asp-Cys-Cys-Ser-Arg-Phe-Xaa₆-Ile-Xaa₅-Xaa₆-Asn-Asp-Phe (SEQ ID NO:12);

Asn-Ala-Cys-Cys-Ile-Val-Arg-Gln-Cys-Cys (SEQ ID NO:13);

Asn-Gly-Cys-Cys-Arg-Ala-Gly-Asp-Cys-Cys-Ser (SEQ ID NO:14);

Cys-Cys-Xaa₁-Arg-Arg-Leu-Ala-Cys-Cys-Ile-Ile (SEQ ID NO:15);

Cys-Cys-Xaa₁-Asn-Xaa₅-Xaa₁-Cys-Cys-Phe-Ile (SEQ ID NO:16);

Gly-Cys-Cys-Ala-Met-Leu-Thr-Cys-Cys-Val (SEQ ID NO:17);

Leu-Cys-Cys-Val-Thr-Xaa₆-Asp-Xaa₃-Cys-Cys-Xaa₆-Xaa₃-Xaa₃ (SEQ ID NO:18); and

Val-Cys-Cys-Arg-Xaa₁-Val-Gln-Asp-Cys-Cys-Ser (SEQ ID NO:19);

wherein Xaa₁ is Pro or hydroxy-Pro; Xaa₂ is Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr or nitro-Tyr; Xaa₃ is Trp or halo-Trp; Xaa₄ is Gln or pyro-Glu; Xaa₅ is Lys, N-methyl-Lys, N,N-dimethyl-Lys or N,n,N-trimethyl-Lys, Xaa₆ is Glu or gamma-carboxy-Glu (Gla); and the C-terminus contains a carboxyl or amide group.

31. The method of claim 30, wherein the pharmaceutical composition is modified to contain an O-glycan, an S-glycan or an N-glycan.